UNITED STATES PATENT AND TRADEMARK OFFICE

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/609,383	07/01/2003	Richard J. Feldmann	3279-Z	4498

7590   01/23/2007

Law Office of Jim Zegeer
Suite 108
801 North Pitt Street
Alexandria, VA 22314

EXAMINER
BRUSCA, JOHN S

ART UNIT	PAPER NUMBER
1631	

SHORTENED STATUTORY PERIOD OF RESPONSE	MAIL DATE	DELIVERY MODE
3 MONTHS	01/23/2007	PAPER

Please find below and/or attached an Office communication concerning this application or proceeding.

If NO period for reply is specified above, the maximum statutory period will apply and will expire 6 MONTHS from the mailing date of this communication.

PTOL-90A (Rev. 10/06)

	Application No.	Applicant(s)
	10/609,383	FELDMANN, RICHARD J.
Office Action Summary	Examiner	Art Unit	
	John S. Brusca	1631	

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --

Period for Reply

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE <u>3</u> MONTH(S) OR THIRTY (30) DAYS, WHICHEVER IS LONGER, FROM THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

Status

1) ☒ Responsive to communication(s) filed on <u>04 December 2006 and 22 September 2006</u>.
2a) ☐ This action is **FINAL**. 2b) ☒ This action is non-final.
3) ☐ Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

Disposition of Claims

4) ☒ Claim(s) <u>*1-12*</u> is/are pending in the application.

    4a) Of the above claim(s) <u>*3-12*</u> is/are withdrawn from consideration.

5) ☐ Claim(s) _____ is/are allowed.
6) ☒ Claim(s) <u>*1 and 2*</u> is/are rejected.
7) ☐ Claim(s) _____ is/are objected to.
8) ☐ Claim(s) _____ are subject to restriction and/or election requirement.

Application Papers

9) ☒ The specification is objected to by the Examiner.
10) ☐ The drawing(s) filed on _____ is/are: a) ☐ accepted or b) ☐ objected to by the Examiner.

    Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).

    Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).

11) ☐ The oath or declaration is objected to by the Examiner. Note the attached Office Action or form PTO-152.

Priority under 35 U.S.C. § 119

12) ☐ Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).

    a) ☐ All b) ☐ Some * c) ☐ None of:

      1. ☐ Certified copies of the priority documents have been received.

      2. ☐ Certified copies of the priority documents have been received in Application No. _____.

      3. ☐ Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).

    * See the attached detailed Office action for a list of the certified copies not received.

Attachment(s)

1) ☒ Notice of References Cited (PTO-892)
2) ☐ Notice of Draftsperson's Patent Drawing Review (PTO-948)
3) ☐ Information Disclosure Statement(s) (PTO/SB/08)
    Paper No(s)/Mail Date _____.

4) ☐ Interview Summary (PTO-413)
    Paper No(s)/Mail Date. _____.
5) ☐ Notice of Informal Patent Application
6) ☒ Other: *Notice to Comply*.

DETAILED ACTION

Priority

1.   The applicant's claim for benefit of Provisional U.S. Application No. 60/393558 is

accepted in view of the amendment to the specification filed 22 September 2006.

Information Disclosure Statement

2.   The information disclosure statement filed 19 November 2003 fails to comply with 37

CFR 1.98(a)(2), which requires a legible copy of each U.S. and foreign patent; each publication

or that portion which caused it to be listed; and all other information or that portion which caused

it to be listed. It has been placed in the application file, but the information referred to therein

has not been considered.

3.   The applicants have filed a CD-ROM with references in lieu of a paper copy, however

foreign and non-patent references must be submitted as paper copies. At this time the Office has

no provisions to accept references on a CD-ROM.

4.   The Information Disclosure Statement filed 19 November 2003 does not contain a paper

copy of each reference listed on the list of references as discussed above. If the applicants

provide a legible copy of the missing references in response to this Office action, the references

will be considered under 37 CFR 1.97(f), and a signed copy of the list of references indicating

consideration of the missing references will be provided to the applicants without the necessity

of the applicants filing a second Information Disclosure Statement.

5.   Applicant's arguments filed 22 September 2006 have been fully considered but they are

not persuasive. The applicants state that a CD-ROM should be considered to be a legible copy,

however the Office at present does not accept CD-ROMs as meeting the requirement of a legible

copy of cited references in an information disclosure statement.

Specification

6.   This application contains sequence disclosures that are encompassed by the definitions

for nucleotide and/or amino acid sequences set forth in 37 CFR §§ 1.821(a)(1) and (a)(2).

However, this application fails to comply with the requirements of 37 CFR §§ 1.821-1.825 for

the following reasons:

SEQ ID NOS have been added to the specification in response to the objection to the

specification in the Office action mailed 22 March 2006, however the computer readable form

associated with the new sequence listing is defective. Please see the Notice to comply and RSL

error report attached to this Office action.

Claim Rejections - 35 USC § 112

7.   The following is a quotation of the first paragraph of 35 U.S.C. 112:

> The specification shall contain a written description of the invention, and of the manner and process of making
> and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it
> pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode
> contemplated by the inventor of carrying out his invention.

8.   Claims 1 and 2 are rejected under 35 U.S.C. 112, first paragraph, as containing subject

matter which was not described in the specification in such a way as to enable one skilled in the

art to which it pertains, or with which it is most nearly connected, to use the invention.

In In re Wands (8 USPQ2d 1400 (CAFC 1988)) the CAFC considered the issue of

enablement in molecular biology. The CAFC summarized eight factors to be considered in a

determination of "undue experimentation." These factors include: (a) the quantity of

experimentation necessary; (b) the amount of direction or guidance presented; (c) the presence or

absence of working examples; (d) the nature of the invention; (e) the state of the prior art; (f) the relative skill of those in the art; (g) the predictability of the art; and (h) the breadth of the claims.

In considering the factors for the instant claims:

a) Quantity of experimentation: The only utility asserted by the specification is to use connectron symmetries to predict control of gene expression (see for example pages 11, 15, and 16 of the specification).In order to practice the claimed invention one of skill in the art must identify and use a connectron to predict regulation of gene expression. In some embodiments changes in connectron behavior that correlate with changes in gene expression is monitored or effected. For the reasons discussed below, there would be an unpredictable amount of experimentation required to practice the claimed invention.

b) The amount of direction or guidance presented: The claimed invention is a method of identification of sequences that have a connectron relationship and act to modulate gene expression. On page 3, the specification defines connectrons as a tetradic structure between two sequences in an RNA transcript of a genomic sequence and two sequences in double stranded genomic DNA. The specification speculates without evidence on page 7 that triple-stranded (triplex) structures will form between RNA and double stranded DNA in chromatin where connectron symmetries are identified. The specification does not provide guidance that there are any limitations on formation of triplex structures, and only implies that regions of RNA with identical sequence to one strand of a double stranded DNA sequence will form triplex structures. The specification does not address why all RNA transcripts of genes would not form a continuous triplex structure with the gene from which it is transcribed. The specification provides guidance to identify connectron symmetries in genomic sequences. The specification

does not provide detailed guidance to use identified connectron symmetries because the specification does not show whether or not connectrons form within cells or have an effect on gene expression. The specification does not provide specific guidance for monitoring or effecting changes in connectron behavior that correlate with gene expression.

c) The presence or absence of working examples: The specification provides working examples of identification of connectron symmetries by computer-mediated searching of genomic sequences. However, the specification does not provide evidence that connectron symmetries in genomic sequences result in formation of triplex RNA-DNA structures or that if connectron triplex structures do exist that connectrons control gene expression. The specification does not provide working examples of using identified connectron symmetries to predict effects on gene expression. The specification does not provide working examples of monitoring or effecting changes in connectron behavior that correlate with gene expression.

d) The nature of the invention: The nature of the invention, gene expression control, is complex.

e) The state of the prior art: One of skill in the art, after reading the specification, would not know that connectron symmetries identified by computer-mediated searches of genomic sequences would allow for prediction of gene expression of genes that have connectron symmetries. The specification does not provide experimental evidence that connectron symmetries cause modulation of gene expression. Neither the prior art nor post-filing art shows connectrons. Mattick (published in 2001, one year after the effective instant filing date) reviews effects of RNA molecules on gene regulation. Mattick does not show connectrons as defined in the instant specification. Chan et al. reviews triplex DNA formation. Chan et al. shows in figures

1A-C that short stretches of oligonucleotides may form parallel or antiparallel triplex structures. Chan et al. shows in figures 1B that parallel triplex forming oligonucleotides form bonds between C and T residues of the oligonucleotide and G and A residues of the double stranded DNA molecule. Figure 1C shows that antiparallel triplex forming oligonucleotides form bonds between A, G, and T residues of the oligonucleotide and A, G, and A residues of the double stranded DNA. Chan et al. characterize the limited range of base pairing possibilities in triplex structures as pyrimidine binding motifs or purine binding motifs. Chan et al. describe on pages 268-273 the unpredictability and difficulty of forming desired triplex structures that are limited to the purine motif or the pyrimidine motif. Chan et al. does not show a mechanism that allows for triplex structures to form with any and all regions of identity between an RNA transcript and a region of double stranded DNA that has an identical sequence in one of the two strands of DNA, as required for connectron formation as defined in the instant specification.

f) The relative skill of those in the art: The skill of those in the art of gene expression is high.

g) The predictability of the art: The predictability of the relationship of connectron symmetries and gene expression is unknown in the prior art and is not described in the instant specification.

h) The breadth of the claims: The claims are broad in that they are drawn to identification and modulation of connectron symmetries whose relationship to gene expression is not established.

The skilled practitioner would first turn to the instant specification for guidance in using the claimed invention. However, the specification lacks any evidence that connectrons form in

cells or that connectron symmetries are related to gene expression. As such, the skilled

practitioner would turn to the prior art for such guidance, however the prior art does not discuss

connectron symmetries. Chan et al. shows that triplex formation occurs only with

oligonucleotides with a purine rich or pyrimidine rich motif, rather than with any identical

sequence as suggested in the specification. Finally, said practitioner would turn to trial and error

experimentation to determine a relationship between connectron symmetries and gene

expression. Such amounts to undue experimentation.

9.   Applicant's arguments filed 22 September 2006 have been fully considered but they are

not persuasive. The applicants state that one of skill in the art would be able to determine

connectron symmetries by analysis of genomic sequence data. However, although the

specification provides guidance and working examples of determining connectron symmetries in

genomic data, the specification does not enable one of skill in the art to use connectron

symmetries to predict regulation of gene expression because the specification does not show that

connectron symmetries have any relationship to triplex formation or control of gene expression

in cells, and the only utility asserted by the specification is to use connectron symmetries to

predict control of gene expression (see for example pages 11, 15, and 16 of the specification).

Conclusion

10.   Any inquiry concerning this communication or earlier communications from the

examiner should be directed to John S. Brusca whose telephone number is 571 272-0714. The

examiner can normally be reached on M-F 8:30 AM - 5:00 PM.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's

supervisor, Andrew Wang can be reached on 571 272-0811. The fax phone number for the

organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent

Application Information Retrieval (PAIR) system. Status information for published applications

may be obtained from either Private PAIR or Public PAIR. Status information for unpublished

applications is available through Private PAIR only. For more information about the PAIR

system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR

system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would

like assistance from a USPTO Customer Service Representative or access to the automated

information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

John S. Brusca
Primary Examiner
Art Unit 1631

jsb

NOTICE TO COMPLY WITH SEQUENCE RULES	Application No. 10/609,383	Applicant(s) FELDMANN, RICHARD J.
	Examiner John S. Brusca	Art Unit 1631

NOTICE TO COMPLY WITH REQUIREMENTS FOR·PATENT APPLICATIONS CONTAINING NUCLEOTIDE SEQUENCE AND/OR AMINO ACID SEQUENCE DISCLOSURES

The nucleotide and/or amino acid sequence disclosure contained in this application does not comply with the requirements for such a disclosure as set forth in 37 CFR 1.821-1.825 for the following reasons:

☒ 1. This application clearly fails to comply with the requirements of 37 CFR 1.821-1.825. Applicant's attention is directed to these regulations, published at 1114 OG 29, May 15, 1990 and at 55 FR 18230, May 1, 1990.

☐ 2. This application does not contain, as a separate part of the disclosure on paper copy, a "Sequence Listing" as required by 37 CFR 1.821(c).

☐ 3. A copy of the "Sequence Listing" in computer readable form has not been submitted as required by 37 CFR 1.821(e).

☒ 4. A copy of the "Sequence Listing in computer readable form has been submitted. However the content of the computer readable form does not comply with the requirements of 37 CFR 1.822 and/or 1.823, as indicated on the attached copy òf the marked up "Raw Sequence Listing".

☐ 5. The computer readable form that has been filed with this application has been found to be damaged and/or unreadable. A Substitute computer readable form must be submitted as required by 37 CFR 1.825(d).

☐ 6. The paper copy of the "Sequence Listing" is not the same as the computer readable form of the "Sequence Listing" as required by 37 CFR 1.821(e).

☐ 7. Other:

Applicant must provide:

☐ An initial or ☒ A substitute computer readable form copy of the Sequence Listing.

☐ An initial or ☐ A Substitute paper copy of the Sequence Listing as well as an amendment directing its entry into the specification.

☐ A statement that the content of the paper and computer readable copies are the same, and, where applicable, include no new matter, as required by 37 CFR 1.821(e), (f), or (g) or 1.825(b) or (d).

FOR QUESTIONS PLEASE CONTACT:

Rules Interpretation (703) 308-4216
CRF Submission Help (703) 308 4212
PatentIn software help (703) 308 6856

PLEASE RETURN A COPY OF THIS NOTICE WITH YOUR RESPONSE

STIC Biotechnology Systems Branch

RAW SEQUENCE LISTING
ERROR REPORT

The Biotechnology Systems Branch of the Scientific and Technical Information Center (STIC) detected errors when processing the following computer readable form:

Application Serial Number: *10/609,383 β*
Source: *,fw/b*
Date Processed by STIC: *9/27/06*

THE ATTACHED PRINTOUT EXPLAINS DETECTED ERRORS.
PLEASE FORWARD THIS INFORMATION TO THE APPLICANT BY EITHER:
1) INCLUDING A COPY OF THIS PRINTOUT IN YOUR NEXT COMMUNICATION TO THE APPLICANT, WITH A NOTICE TO COMPLY or,
2) TELEPHONING APPLICANT AND FAXING A COPY OF THIS PRINTOUT, WITH A NOTICE TO COMPLY
FOR CRF SUBMISSION AND PATENTIN SOFTWARE QUESTIONS, PLEASE CONTACT
MARK SPENCER, TELEPHONE: 571-272-2510; FAX: 571-273-0221

TO REDUCE ERRORED SEQUENCE LISTINGS, PLEASE USE THE <u>CHECKER VERSION 4.4.0 PROGRAM</u>, ACCESSIBLE THROUGH THE U.S. PATENT AND TRADEMARK OFFICE WEBSITE. SEE BELOW FOR ADDRESS:
http://www.uspto.gov/web/offices/pac/checker/chkrnote.htm

Applicants submitting genetic sequence information electronically on diskette or CD-Rom should be aware that there is a possibility that the disk/CD-Rom may have been affected by treatment given to all incoming mail.
Please consider using alternate methods of submission for the disk/CD-Rom or replacement disk/CD-Rom.
Any reply including a sequence listing in electronic form should NOT be sent to the 20231 zip code address for the United States Patent and Trademark Office, and instead should be sent via the following to the indicated addresses:

1. EFS-Bio (<http://www.uspto.gov/ebc/efs/downloads/documents.htm>, EFS Submission User Manual - ePAVE)
2. U.S. Postal Service: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450
3. Hand Carry, Federal Express, United Parcel Service, or other delivery service (EFFECTIVE 01/14/05):
   U.S. Patent and Trademark Office, Mail Stop Sequence, Customer Window, Randolph Building, 401 Dulany Street, Alexandria, VA 22314

Revised 01/10/06

Raw Sequence Listing Error Summary

ERROR DETECTED	SUGGESTED CORRECTION	SERIAL NUMBER: _10/609,383B_

ATTN: NEW RULES CASES: PLEASE DISREGARD ENGLISH "ALPHA" HEADERS, WHICH WERE INSERTED BY PTO SOFTWARE

1. ____Wrapped Nucleics Wrapped Aminos — The number/text at the end of each line "wrapped" down to the next line. This may occur if your file was retrieved in a word processor after creating it. Please adjust your right margin to .3; this will prevent "wrapping."

2. ____Invalid Line Length — The rules require that a line not exceed 72 characters in length. This includes white spaces.

3. ____Misaligned Amino Numbering — The numbering under each 5th amino acid is misaligned. Do not use tab codes between numbers; use space characters, instead.

4. ____Non-ASCII — The submitted file was not saved in ASCII(DOS) text, as required by the Sequence Rules. Please ensure your subsequent submission is saved in ASCII text.

5. ____Variable Length — Sequence(s)____contain n's or Xaa's representing more than one residue. Per Sequence Rules, each n or Xaa can only represent a single residue. Please present the maximum number of each residue having variable length and indicate in the <220>-<223> section that some may be missing.

6. ____PatentIn 2.0 "bug" — A "bug" in PatentIn version 2.0 has caused the <220>-<223> section to be missing from amino acid sequences(s) _____. Normally, PatentIn would automatically generate this section from the previously coded nucleic acid sequence. Please manually copy the relevant <220>-<223> section to the subsequent amino acid sequence. **This applies to the mandatory <220>-<223> sections for Artificial or Unknown sequences.**

7. ____Skipped Sequences (OLD RULES) — Sequence(s) _____ missing. If intentional, please insert the following lines for each skipped sequence:
   (2) INFORMATION FOR SEQ ID NO:X: (insert SEQ ID NO where "X" is shown)
   (i)   SEQUENCE CHARACTERISTICS: (Do not insert any subheadings under this heading)
   (xi) SEQUENCE DESCRIPTION:SEQ ID NO:X: (insert SEQ ID NO where "X" is shown)
   This sequence is intentionally skipped
   Please also adjust the "(ii) NUMBER OF SEQUENCES:" response to include the skipped sequences.

8. ____Skipped Sequences (NEW RULES) — Sequence(s) _____ missing. If intentional, please insert the following lines for each skipped sequence.
   <210> sequence id number
   <400> sequence id number
   000

9. ____Use of n's or Xaa's (NEW RULES) — Use of n's and/or Xaa's have been detected in the Sequence Listing.
   Per 1.823 of Sequence Rules, use of <220>-<223> is MANDATORY if n's or Xaa's are present.
   In <220> to <223> section, please explain location of n or Xaa, and which residue n or Xaa represents.

10. ____Invalid <213> Response — Per 1.823 of Sequence Rules, the only valid <213> responses are: Unknown, Artificial Sequence, or scientific name (Genus/species). <220>-<223> section is required when <213> response is Unknown or is Artificial Sequence. (see item 11 below)

11. ____Use of <220> — Sequence(s) _____ missing the <220> "Feature" and associated numeric identifiers and responses. Use of <220> to <223> is MANDATORY if <213> "Organism" response is "Artificial Sequence" or "Unknown." Please explain source of genetic material in <220> to <223> section or use "chemically synthesized" as explanation. (See "Federal Register," 06/01/1998, Vol. 63, No. 104, pp. 29631-32), also Sec. 1.823 of Sequence Rules

12. ____PatentIn 2.0 "bug" — Please do not use "Copy to Disk" function of PatentIn version 2.0. This causes a corrupted file, resulting in missing mandatory numeric identifiers and responses (as indicated on raw sequence listing). Instead, please use "File Manager" or any other manual means to copy file to floppy disk.

13. ____Misuse of n/Xaa — "n" can only represent a single nucleotide; "Xaa" can only represent a single amino acid

AMC – STIC Systems Branch – 03/02/06
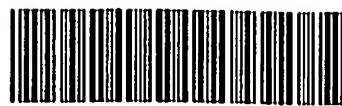
IFW16

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RAW SEQUENCE LISTING                    DATE: 09/27/2006
PATENT APPLICATION:  US/10/609,383B     TIME: 15:13:28

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       Output Set: N:\CRF4\09272006\J609383B.raw
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(handwritten: see pp 7-8, too)

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      1 <110> APPLICANT: Feldmann, Richard J.; Connectron Holding, Inc.
      3 <120> TITLE OF INVENTION: Synthetic Connectron
      5 <130> FILE REFERENCE: Jim Zegeer Law Offices - 703-684-8333
C-->  7 <140> CURRENT APPLICATION NUMBER: US/10/609,383B
C-->  7 <141> CURRENT FILING DATE: 2003-07-01
      8 <150> PRIOR APPLICATION NUMBER: US 60/393,558 and US 09/866,925
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     19 <213> ORGANISM: Saccharomyces cerevisiae complete genome.
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     28 ggtggtaatg  atgaagtaat  ttcctgactt  gttgttgtac  tggtaacagg  tggtaatgaa    120
     29 gaagtaattt  cctgacttgt  tgttgcactg  gtaacaggtg  gtaatgatga  agtaatttcc    180
     30 tgacttgttg  ttgtactggt  aacaggtggt  aatgatg                                217
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     46 ggtggtaatg  atgaagtaat  ttcctgactt  gttgttgtac  tggtaacagg  tggtaatgaa    120
     47 gaagtaattt  cctgacttgt  tgttgcactg  gtaacaggtg  gtaatgatga  agtaatttcc    180
     48 tgacttgttg  ttgtactggt  aacaggtggt  aatgatgaag  cagtttcctg  gcttgt        236
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     65 ctttggttac  cgtgacatcc  tgcgaatctc  atgtgtgcac  tgaatc                     166
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(handwritten annotations: "Does Not Comply Corrected Diskette Needed", "correct format", "list as <150> 60/393,558 <151> 2002-07-05, followed by <150> 09/866,925 <151> 2001-05-30 on line below")

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PATENT APPLICATION: US/10/609,383B   TIME: 15:13:28

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          Output Set: N:\CRF4\09272006\J609383B.raw

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   74 <220> FEATURE:
   75 <222> LOCATION: (4626130)...(4626166)
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975
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  132 <213> ORGANISM: Escherichia coli K-12 MG1655 complete genome.
  134 <220> FEATURE:
  135 <222> LOCATION: (698713)...(698766)
  136 <223> OTHER INFORMATION: Chromosome = 1 Strand = negative ConnectronObjectNumber =
809
  139 <400> SEQUENCE: 8
  140 tctgatgaca aacgccaaac tgcctgatgc gctacgctta tcaggcctac gcag   54
  143 <210> SEQ ID NO: 9
  145 <211> LENGTH: 36
  146 <212> TYPE: DNA

147 <213> ORGANISM: Escherichia coli K-12 MG1655 complete genome.

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Input Set : P:\synthetic.txt
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```
     149 <220> FEATURE:
     150 <222> LOCATION: (757718)...(757753)
     151 <223> OTHER INFORMATION: Chromosome = 1 Strand = negative ConnectronObjectNumber =
975
     154 <400> SEQUENCE: 9
     155 ttacgcctga tgcgctgcgc ttatcaggcc tacggg                                  36
     158 <210> SEQ ID NO: 10
     160 <211> LENGTH: 16
     161 <212> TYPE: DNA
     162 <213> ORGANISM: Saccharomyces cerevisiae complete genome - problem.
     164 <220> FEATURE:
     165 <222> LOCATION: (221330)...(221345)
     166 <223> OTHER INFORMATION: Chromosome = 2 Strand = positive ConnectronObjectNumber =
792a
     169 <400> SEQUENCE: 10
     170 tatatatatg tcactg                                                         16
     173 <210> SEQ ID NO: 11
     175 <211> LENGTH: 16
     176 <212> TYPE: DNA
     177 <213> ORGANISM: Saccharomyces cerevisiae complete genome - problem.
     179 <220> FEATURE:
     180 <222> LOCATION: (221346)...(221361)
     181 <223> OTHER INFORMATION: Chromosome = 2 Strand = positive ConnectronObjectNumber =
793
     184 <400> SEQUENCE: 11
     185 tattgcatgc tggatg                                                         16
     188 <210> SEQ ID NO: 12
     190 <211> LENGTH: 539
     191 <212> TYPE: DNA
     192 <213> ORGANISM: Saccharomyces cerevisiae complete genome - problem.
     194 <220> FEATURE:
     195 <222> LOCATION: (448454)...(448992)
     196 <223> OTHER INFORMATION: Chromosome = 5 Strand = positive ConnectronObjectNumber =
4749
     199 <400> SEQUENCE: 12
     200 tatatatatg tcactgtatt gcatgctgga tggtgttaga caaggccgta gggacatata        60
     201 gcatctagga agtaaccttg tacgaaaata ggcaatattt cctgtttagg cgattgtgac       120
     202 gcagatttta gtccaacgat ctagcgtcaa ggaatttttt tatagtggga cattgcacca       180
     203 aggaagtaac ttgatacgtc gtgggtgaat gggtctgttt tcttattcgg cggggtaata       240
     204 catttttggg ggaagtttgt ctgtctgacg cgccatatgt aggtacgcca aaaagggctc       300
     205 ctctacttcg aagcgcgagg tcgtatacct aataaggaaa tgtaatttat aactttttat       360
     206 tatattggtc ttttcgagag cggaacgtag gtccatgttt aaagtatcca agagaatatc       420
     207 cacgaagcgg ctgagcaacg aacagaatcc tggttctcct cgactaagca gatagttaag       480
     208 atactgtgca ccatggaaat tgaaaacgaa agtacgtacc gactacttta tttttgcag        539
     210 <210> SEQ ID NO: 13
     212 <211> LENGTH: 158
     213 <212> TYPE: DNA
     214 <213> ORGANISM: Saccharomyces cerevisiae complete genome - problem.
     216 <220> FEATURE:
     217 <222> LOCATION: (24863)...(25028)
     218 <223> OTHER INFORMATION: Chromosome = 5 Strand = negative ConnectronObjectNumber =
4824a
     221 <400> SEQUENCE: 13
```

222 tatatatatg tcactgtatt gcatgctgga tggtgttaga caaggccgta gggacatata   60

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Input Set : P:\synthetic.txt
Output Set: N:\CRF4\09272006\J609383B.raw

```
223 gcatctagga  agtaaccttg  tacgaaaata  ggcaatattt  cctgtttagg  cgattgtgac          120
224 gcagatttta  gtccaacgat  ctagcgtcaa  ggaatttt                                    158
226 <210> SEQ ID NO: 14
228 <211> LENGTH: 134
229 <212> TYPE: DNA
230 <213> ORGANISM: Halobacterium sp. NRC-1 complete genome.
232 <220> FEATURE:
233 <222> LOCATION: (732401)...(732534)
234 <223> OTHER INFORMATION: Chromosome = 1 Strand = positive ConnectronObjectNumber =
6612
237 <400> SEQUENCE: 14
238 ttcatcacag  acgaggacga  gcgcggccaa  gtggggatcg  gcacactcat  cgtgttcatc           60
239 gcgatggtgc  tggtcgccgc  gatcgccgcc  ggcgtcctca  tcaacactgc  cggctacctc          120
240 caatccaagg  ggtc                                                                134
243 <210> SEQ ID NO: 15
245 <211> LENGTH: 193
246 <212> TYPE: DNA
247 <213> ORGANISM: Halobacterium sp. NRC-1 complete genome.
249 <220> FEATURE:
250 <222> LOCATION: (733018)...(733209)
251 <223> OTHER INFORMATION: Chromosome = 1 Strand = positive ConnectronObjectNumber =
6644a
254 <400> SEQUENCE: 15
255 gacgagcgcg  gtcaagtggg  gatcggcaca  ctcatcgtgt  tcatcgcgat  ggtgctggtc           60
256 gccgcgatcg  ccgccggcgt  cctcatcaac  accgccggct  acctccaatc  caaggggtcg          120
257 gcaaccggtg  aggaagcctc  cgcacaggtc  tccaaccgca  tcaacatcgt  ctccgcgtac          180
258 ggcaacgtca  aca                                                                 193
261 <210> SEQ ID NO: 16
263 <211> LENGTH: 85
264 <212> TYPE: DNA
265 <213> ORGANISM: Halobacterium sp. NRC-1 complete genome.
267 <220> FEATURE:
268 <222> LOCATION: (773399)...(773483)
269 <223> OTHER INFORMATION: Chromosome = 1 Strand = positive ConnectronObjectNumber =
6852
272 <400> SEQUENCE: 16
273 gtggggatcg  gcacgctcat  cgtgttcatc  gcgatggtgc  tggtcgccgc  gatcgccgcc           60
274 ggcgtcctca  tcaacactgc  cggct                                                    85
277 <210> SEQ ID NO: 17
279 <211> LENGTH: 121
280 <212> TYPE: DNA
281 <213> ORGANISM: Pseudomonas aeruginosa PA01, complete genome.
283 <220> FEATURE:
284 <222> LOCATION: (4832718)...(4832838)
285 <223> OTHER INFORMATION: Chromosome = 1 Strand = positive ConnectronObjectNumber =
53464
288 <400> SEQUENCE: 17
289 gccaacatcg  aggccctcaa  cagccgcacg  gtgaacatcg  gccagatcct  cgaagtgatc           60
290 aagggcatct  ccgagcagac  caacctgctc  gccctcaacg  ccgccatcga  agccgcgcgc          120
291 g                                                                              121
294 <210> SEQ ID NO: 18
296 <211> LENGTH: 194
297 <212> TYPE: DNA
```

298 <213> ORGANISM: Pseudomonas aeruginosa PA01, complete genome.
300 <220> FEATURE:
301 <222> LOCATION: (4836528)...(4836720)
302 <223> OTHER INFORMATION: Chromosome = 1 Strand = positive ConnectronObjectNumber =
53531
305 <400> SEQUENCE: 18
306 ggacggcaaa caggtggtcg agcagaccat ccgcgcgatg aacgagcttt ccgagaagat   60
307 cagcgcctcc tgcgccaaca tcgaggccct caacagccgc acggtgaaca tcggccagat   120
308 cctcgaagtg atcaagggca tctccgagca gaccaacctg ctcgccctca acgccgccat   180
309 cgaagccgcg cgcg   194
312 <210> SEQ ID NO: 19
314 <211> LENGTH: 169
315 <212> TYPE: DNA
316 <213> ORGANISM: Pseudomonas aeruginosa PA01, complete genome.
318 <220> FEATURE:
319 <222> LOCATION: (4838678)...(4838846)
320 <223> OTHER INFORMATION: Chromosome = 1 Strand = positive ConnectronObjectNumber =
53549a
323 <400> SEQUENCE: 19
324 accatccgcg cgatgaacga gctttccgag aagatcagcg cctcctgcgc caacatcgag   60
325 gccctcaaca gccgcacggt gaacatcggc cagatcctcg aagtgatcaa gggcatctcc   120
326 gagcagacca acctgctcgc cctcaacgcc gccatcgaag ccgcgcgcg   169
329 <210> SEQ ID NO: 20
331 <211> LENGTH: 36
332 <212> TYPE: DNA
333 <213> ORGANISM: Sequence Recognized by Synthetic DNA Binding Protein.
335 <220> FEATURE:
338 <400> SEQUENCE: 20
339 tccccatgag catagatatg caggtaggcg gcaagt
342 <210> SEQ ID NO: 21
344 <211> LENGTH: 136
345 <212> TYPE: DNA
346 <213> ORGANISM: Vibrio cholerae chromosome I, complete chromosome.
348 <220> FEATURE:
349 <222> LOCATION: (952641)...(952777)
350 <223> OTHER INFORMATION: Chromosome = 1 Strand = negative ConnectronObjectNumber =
607
353 <400> SEQUENCE: 21
354 tgtatatacc caaactactt ggagttgcag gtaggcggca agtgagtgag tccccatgag   60
355 catagataga ctatgtgatt ggggtgaacg aacgtagcca acaccgctgc agcttcaagt   120
356 aggaagggta tacctt   136
359 <210> SEQ ID NO: 22
361 <211> LENGTH: 117
362 <212> TYPE: DNA
363 <213> ORGANISM: Vibrio cholerae chromosome I, complete chromosome.
365 <220> FEATURE:
366 <222> LOCATION: (1005810)...(1005926)
367 <223> OTHER INFORMATION: Chromosome = 1 Strand = negative ConnectronObjectNumber =
646
370 <400> SEQUENCE: 22
371 taccaaaact acttggagtt gcaggtaggc ggcaagagag tgaatcccca tcagcataga   60
372 cagactatgt gattggggtg aacgaacgta gccaataccg ctgcagcttc aagtagg   117
375 <210> SEQ ID NO: 23

[handwritten annotations in right margin:] Invalid response. See item 10 on Error Summary sheet. also, if this is an artificial sequence, please give source of genetic material

[handwritten:] 36

RAW SEQUENCE LISTING ERROR SUMMARY   DATE: 09/27/2006
PATENT APPLICATION: US/10/609,383B   TIME: 15:13:29

Input Set : P:\synthetic.txt
Output Set: N:\CRF4\09272006\J609383B.raw

Invalid Line Length:
The rules require that a line not exceed 72 characters in length. This includes spaces.

```
Seq#:1;  Line(s)  27,28,29,30
Seq#:2;  Line(s)  45,46,47,48
Seq#:3;  Line(s)  63,64,65
Seq#:4;  Line(s)  76,80
Seq#:5;  Line(s)  95
Seq#:6;  Line(s)  110
Seq#:7;  Line(s)  121,125
Seq#:8;  Line(s)  140
Seq#:9;  Line(s)  155
Seq#:10; Line(s)  170
Seq#:12; Line(s)  200,201,202,203,204,205,206,207,208
Seq#:13; Line(s)  222,223,224
Seq#:14; Line(s)  238,239,240
Seq#:15; Line(s)  255,256,257,258
Seq#:16; Line(s)  273,274
Seq#:17; Line(s)  289,290,291
Seq#:18; Line(s)  306,307,308,309
Seq#:19; Line(s)  324,325,326
Seq#:20; Line(s)  339
Seq#:21; Line(s)  354,355,356
Seq#:22; Line(s)  371,372
Seq#:23; Line(s)  385
Seq#:24; Line(s)  400,401,402
Seq#:25; Line(s)  417,418
Seq#:26; Line(s). 425,431
Seq#:27; Line(s)  438,444
Seq#:28; Line(s)  459,460
Seq#:29; Line(s)  475,476,477
Seq#:30; Line(s)  489
Seq#:31; Line(s)  500
Seq#:32; Line(s)  511
Seq#:33; Line(s)  522
Seq#:34; Line(s)  533
```

VERIFICATION SUMMARY   DATE: 09/27/2006
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Input Set : **F:\synthetic.txt**
Output Set: **N:\CRF4\09272006\J609383B.raw**

L:7 M:270 C: Current Application Number differs, Missing <140> CURRENT APPLICATION NUMBER: is Added.
L:7 M:271 C: Current Filing Date differs, Replaced Current Filing Date